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Technical Field 3 The present invention relates to fluid storage 5 apparatus, in particular, but not exclusively, to apparatus for the disruption and storage of cellular 7 fluids. 9 10 Background 11 A knowledge of the constituent components of the 12 cells of cellular fluids, such as deoxyribonucleic 13 acid, are of great importance to the understanding 14 15 of how such cells function. In order to analyse these components from the cells it is necessary to 16 17 cause disruption of the cells. This basically means that the walls of the cells are broken down, thus 18 allowing the constituent components to be removed 19 20 for analysis.

Fluid Storage Apparatus

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Disrupted cellular fluids, that is, cellular fluids 1 2 in which the cell walls have burst, are conventionally stored in a pre-sterilised sealed 3 container which may be further stored in a plastic bag and refrigerated prior to use. 5 disrupted cellular fluids in this manner however, means taking a sample of the disrupted cellular 7 fluid and placing it in the container. 9 handling of the sample greatly increases the risk of contamination and degradation of the sample. 10 Furthermore, the container used to store the sample 11 is often sterilised for re-use, which is expensive 12 13 and further increases the risk of contamination and 14 degradation of the sample. 15 16 It is an object of the present invention to provide a fluid apparatus for the disruption and storage of 17 cellular fluids which obviates or mitigates one or 18 19 more of the disadvantages referred to above. 20 21 Summary of Invention 22 23 According to a first aspect of the present invention 24 there is provided a fluid storage apparatus 25 comprising a first container having a first chamber 26 capable of being filled with a fluid, a second 27 container having a second chamber adapted to receive fluid from said first chamber, the second container 28

having a piston means slideably receivable within

on insertion of said piston means into said first

said first chamber of said first container, wherein,

1 chamber of said first container, fluid is displaced from said first chamber to said second chamber. 3 Preferably the piston means and the second container 4 are integrally formed. 7 Preferably the piston means has a bore which fluidly 8 communicates with the first and second chambers. Preferably the bore has a first portion having a 10 11 first diameter, and a second portion having a second 12 diameter which is smaller than the first diameter. 13 Preferably the first portion of the bore is adjacent 14 the second chamber and the second portion of the 15 bore is remote from the second chamber. 16 17 Preferably the fluid storage apparatus further 18 comprises a sealing means adapted to seal the first 19 20 and second containers together. 21 Preferably the first and second containers are 22 23 adapted to seal together as the fluid is displaced to the second chamber. 24 25 26 Preferably at least one portion of the second chamber is adapted to allow fluid to be removed 27 therefrom. 28 29 30 Preferably the fluid storage apparatus further

includes cutting means adapted to remove a part of

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the apparatus such that the stored fluid may be 1 removed from the second chamber. 2 3 Preferably the fluid storage apparatus is 4 disposable. 6 7 Brief Description of the Drawings 8 Embodiments of the present invention will now be 9 described, by way of example only, with reference to 10 11 the accompanying drawings, in which:-12 13 Fig. 1 is a side view of a first embodiment of a fluid storage apparatus in an initial position, 14 Fig. 2 is a side view of the fluid storage apparatus 15 of Fig. 1 in a storage position, 16 Fig. 3 is a side view of a second embodiment of a 17 fluid storage apparatus in an initial position, and 18 Fig. 4 is a side view of the fluid storage apparatus 19 20 of Fig. 3 in a storage position. 21 Detailed Description 22 23 Referring to Figs. 1 and 2 of the drawings, a fluid 24 25 storage apparatus 10 comprises a first container 12 having a first chamber 14 capable of being filled 26 27 with a volume of cellular fluid, and a second container 16 having a second chamber 18 and a piston 28 A cellular fluid is considered here as being a 29

fluid which is comprised of a large number of cells. For example biological or man-made materials, such

as blood, tissue homogenate and saliva.

· 1

2 The piston 20 and the second container 16 are

3 integrally formed.

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5. The piston 20 has a central bore 26 which allows

fluid communication between the first and second

7 chambers 14 and 18 when in use.

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9 The first container 12 is substantially cylindrical

10 and defines the first chamber 14, which has a first

11 portion 22 which is also substantially cylindrical,

12 and a second portion 24 located adjacent the first

13 portion 22 which is semi-spherical.

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15 The second container 16 is again substantially

16 cylindrical and defines the second chamber 18 which

is also substantially cylindrical. The second

18 chamber 18 is adapted to store the cellular fluid

19 when the apparatus 10 is in use.

20 21

The second container 16 also comprises a piston 20

22 which extends in a longitudinal direction from the

23 second chamber 18. The piston 20 has a central bore

24 26. The bore 26 has a first portion 26a adjacent

25 the second chamber 18 and a second portion 26b

26 remote from the second chamber 18. The first

27 portion 26a has a first diameter and the second

28 portion 26b has a second diameter which is smaller

29 than the first diameter. The piston 20 is slidably

30 engageable with the first portion 22 of the first

31 container 12. The piston 20 and the first portion

32 22 are sized such that, when they are engaged with

1 one another, a seal is formed therebetween by virtue an interference fit created between the side of the piston 20 and the side of the first chamber 14. 3 interference fit is considered here as meaning a 4 fixed connection between two components which arises 5 б by virtue of friction between the two components. Thus, once the piston 20 is at least partially inserted in the first chamber 14, the first chamber 8 14, the second chamber 18 and the central bore 26 define a sealed volume, which prevents the 10 surrounding air from contaminating or degrading the 11 12 fluid in the apparatus 10. 13 14 The second chamber 18 may have a portion (not shown) which is adapted to allow fluid to be removed 15 16 therefrom. For example, the second chamber 18 may have a thinner wall portion which would allow the 17 insertion of a syringe for extraction of the fluid. 18 19 The typical volume of sample contained within the 20 fluid storage apparatus 10 is approximately 5 ml, 21 although other volumes may be used. 22 23 Prior to use the first and second containers 12 and 24 25 16 are sterilised. 26 In operation, the first chamber 14 of the first 27 container 12 is filled with a sample of cellular 28 The piston 20 is then inserted into the 29 first portion 22 of the first chamber 14 and the 30 first and second containers 12 and 16 are then urged 31

together by means of applying longitudinal forces A 1 and B to their respective end portions 28 and 30. 2 3 The first and second containers 12 and 16 are 4 brought together by a machine (not shown) which 5 applies the requisite amount of force to the end 6 portions 28 and 30. 7 8 As the first and second containers 12 and 16 are 9 brought together the cellular fluid contained in the 10 first chamber 14 is forced by the piston 20 through 11 the central bore 26 and into the second chamber 18. 12 Due to the sealing fit of the piston 20 and the 13 first chamber 14, no fluid can escape between the 14 piston 20 and the first chamber 14. 15 16 The longitudinal forces A and B are applied to the 17 end portions 28 and 30 of the first and second 18 containers 12 and 16 until all the cellular fluid 19 has been transferred from the first chamber 14 to 20 the second chamber 18. Typically, the first and 21 second containers 12 and 16 are brought together in 22 less than 1 millisecond. 23 24 The process of bringing the first and second 25 containers 12 and 16 together in the manner 26 described above causes the cells of the cellular 27 fluid to be disrupted. By forcing the piston 20 28 into the first portion 22 of the first chamber 14, 29 the cellular fluid contained within the first 30 31 chamber 14 is pressurised and is forced through the central bore 26 and into the second chamber 18.

- 1 pressure required to disrupt the cellular fluid is
- 2 dependent upon the type of cellular fluid, but a
- 3 typical pressure is in the region of 40 kpsi (276
- 4 MPa).

- 6 The differing diameters of the first and second
- portions 26a and 26b of the central bore 26 of the
- 8 piston 20 creates a step which aids in the
- 9 disruption of the cellular fluid.

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- 11 The shape, size and configuration of the central
- 12 bore 26 may also be varied depending on the type of
- 13 cellular fluid which is being stored.

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- 15 The cells in the cellular fluid are disrupted by the
- 16 following mechanisms: (a) the boundary level cells
- 17 rupture due to the friction created at the wall of
- 18 the central bore 26 as the fluid passes through the
- 19 central bore 26, (b) the cell walls burst due to the
- 20 pressurisation of the fluid through the central bore
- 21 26, (c) the cells explode as they enter the second
- 22 chamber 18 due to the decrease in pressure and (d)
- 23 the outer cells burst on impact against the inner
- 24 wall of the end portion 30 of the second container
- 25 16.

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- Once the first and second containers 12 and 16 have
- 28 been brought together under the great pressure
- 29 applied, a seal is formed between the piston 20 and
- 30 the first portion 22 of the first chamber 14 by
- 31 virtue of the interference fit described above.
- 32 This seal allows the disrupted cellular fluid sample

to be stored safely and prevents degradation or contamination of the sample.

3

When the disrupted cellular fluid is to be analysed,

5 a syringe, or the like, is inserted through adapted

6 wall portion (not shown) and the fluid is removed.

7 Alternatively, the fluid storage apparatus 10 may

8 further include a cutting means (not shown) which

9 may be used to simply cut open the apparatus 10,

thus allowing the fluid to be removed. The fluid

11 storage apparatus 10 is then disposed of, thus

12 avoiding the need for re-sterilisation.

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14 The preferred material of construction of the fluid

15 storage apparatus 10 is plastic. The first and

16 second container 12 and 16 can be formed by any

17 suitable means, such as injection moulding, for

18 example. The second container 16 and the piston 20

19 are preferably moulded as one piece.

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21 Figs. 3 and 4 of the drawings illustrate a second

22 embodiment of the present invention. Corresponding

23 similar features between the first embodiment and

24 the second embodiment have not been described,

25 although the same reference numerals have been used,

26 prefixed by the number 1.

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28 The second container 116 also comprises a piston 120

29 which extends in a longitudinal direction from the

30 second chamber 118. The piston 120 has a central

31 bore 126. The bore 126 has a first portion 126a

32 adjacent the second chamber 118 and a second portion

# 13/ 2

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126b remote from the second chamber 118. The piston 1 120 also has an orifice 126c at a far end of the 2 piston 120. 5 The first portion 126a has a first diameter and the second portion 126b has a second diameter. first diameter is larger than the second diameter. 8 The piston 120 is slidably engageable with the first 9 portion 122 of the first container 112. The piston 10 120 has ridged sections 121 along its outer surface 11 The piston 120 and the first portion 122 are 12 sized such that, when they are engaged with one 13 another, a seal is formed therebetween by virtue an 14 interference fit created between the ridged sections 15 121 of the piston 120 and the side of the first 16 17 chamber 114. 18 The fluid storage apparatus 110 is operated in the 19 same manner as in the first embodiment. 20 21 When the disrupted cellular fluid is to be analysed, 22 the fluid may be removed from the apparatus 110 by 23 inserting a syringe into a syringe needle access 24 point 135 located adjacent the first chamber 114. 25 26 The fluid storage apparatus 10, 110 therefore 27 obviates or mitigates the disadvantages of previous 28 proposals by providing a fluid storage device which 29 allows the cellular fluid sample to be disrupted as 30 part of the sealing of the apparatus. The apparatus 31

both disrupts the cells of the fluid and stores the

fluid, thereby obviating the need for separate 1 2 disruption and storage. The fluid storage apparatus 10, 110 therefore avoids any contamination or degradation of the cellular sample that 4 5 conventionally arises from the handling of a predisturbed sample. Since the fluid storage apparatus 6 10, 110 is disposable, it also avoids the need for 7 sterilisation after use, which is expensive and further increases contamination and degradation. 9 10 The fluid storage apparatus 10, 110 may, for 11 example, be used is the following procedures: cell 12 disruption, cell rupture, homogenisation, French 13 Press principle, single cell isolation, particle 14 size distribution, emulsifying and cell dispersion 15 of micro-organisms, human and animal tissues organs 16 and fluids, plant and soil. The fluid storage 17 apparatus 10, 110 may also be used, for example, in 18 the following applications: release, extraction and 19 isolation of intracellular organelles and including 20 cytoplasmic and membrane proteins and enzymes, 21 inclusion bodies and isolation, shearing and 22 splicing of deoxyribonucleic acids; and diagnosis of 23 microbial based diseases whereby one of the above 24 procedures is required. 25 26 Modifications and improvements may be made to the 27 above without departing from the scope of the 28 present invention. For example, although the fluid 29 storage apparatus 10, 110 is described as being used 30 with a cellular fluid, it should be appreciated that 31 the fluid storage apparatus 10, 110 could be used 32

1 with any biological or man-made material. although the central bore 26, 126 is shown to be 2 made up of stepped diameter sections 26a, 26b, 126a, 3 126b and 126c, it should be appreciated that the central bore 26, 126 could be shaped in an 5 alternative arrangement. For example the bore could 6 be shaped to form a venturi section. Furthermore, 7 · 8 although the disruption of the cellular fluid is described above as occurring from the pressurising 9 of the fluid through a single central bore 26, 126, 10 11 it should be appreciated that the disruption of the cellular fluid could occur by any type of orifice, 12 13 or orifices. Also, although the typical volume of cellular fluid sample contained within the fluid 14 storage apparatus is described above as being 5 ml, 15 it should be appreciated that the fluid storage 16 apparatus 10, 110 could be adapted to contain any 17 volume of sample. Furthermore, although the fluid 18 storage apparatus 10, 110 is described above as 19 20 being constructed from plastic, it should be 21 appreciated that the fluid storage apparatus 10, 110 22 could be made from alternative materials, including metals such as steel or copper. Also, although the 23 sealing of the fluid storage apparatus 10, 110 is 24 25 described above as the result of an interference fit between the first portion 22, 122 of the first 26 27 chamber 14, 114 and the piston 20, 120, it should be 28 appreciated that the fluid storage apparatus 10, 110 29 could be sealed by any suitable mechanical means. For example, the apparatus 10, 110 could be sealed 30 by clamping the first and second containers 12, 112, 31 32 16, 116 together.

Furthermore, although the removal of the sample from 2 the fluid storage apparatus 10, 110 has been 3 described above as by means of access through a portion of the second chamber 18, 118, it should be 5 appreciated that the sample could be removed from 7 the apparatus 10, 110 by providing a sealed screw cap or a bayonet cap or the like at the end portion 8 30, 130 of the second container 16, 116. 9 Alternatively, the sample could be removed from the 10 apparatus 10, 110 by providing a frangible diaphragm 11 or the like on a wall of the second container 16, 12 116 that allows access to the sample once pierced. 13 Also, the sample could be removed from the apparatus 14 10, 110 by providing a hinged cap (flip-lid) or the 15 like on the second container 16, 116 that could be 16 swung open to allow access to the sample. 17 18 sample could also be removed from the apparatus 10, 110 by providing on the second container 16, 116 a 19 plug or the like which could be pierced by a syringe 20 The sample could also be removed from 21 or the like. the apparatus 10, 110 by providing on the second 22 container 16, 116 a check valve which comprises a 23 24 sealing ball or the like which may be dislodged by a 25 syringe or the like when the sample is removed. sample could also be removed from the apparatus 10, 26 27 110 by providing on the second container 16, 116 a weak portion which may be pierced by a syringe or 28 The sample could also be removed from the the like. 29 30 apparatus 10, 110 by providing on the second container 16, 116 weak sections formed by grooves on 31 the body of the second container 16, 116 (either 32

internal or external) or the like which may be 1 'popped' or 'snapped' out of place allow access to The sample could also be removed from the sample. 3 the apparatus 10, 110 by providing on the second 4 container 16, 116 a breakable spigot or the like on 5 the body of the second container 16, 116 which may 6 be 'snapped' off to allow access to the sample. 7 sample could also be removed from the apparatus 10, 8 110 by providing on then end portion of the piston 9 10 20, 120 a breakable nozzle or the like which may be 'snapped' off to allow access to the sample. 11 12 sample could also be removed from the apparatus 10, 110 by providing a drain device or the like which 13 may be inserted into the end portion 30, 130 of the 14 second container 16, 116. The sample could also be 15 removed from the apparatus 10, 110 by providing on 16 the second container 16, 116 an external tear-off 17 strip or the like which may, for example, be formed 18 19 around the circumference of the second container 16, The external strip is then torn around the 20 circumference of the second container to allow 21 access to the sample. Alternatively, the tear-off 22 strip may be torn by a relative twisting motion 23 between the strip and the container 16, 116. 24 the tear-off strip may be torn-off by providing a 25 26 key device or the like which links with the tear-off strip allowing the strip to be removed upon a 27 turning action of the key. The external tear-off 28 strip may also include a sealing member provide 29 30 between the strip and the container 16, 116. 31 The sample could also be removed from the apparatus 10, 110 by providing on the second container 16, 116 32

a spin weld weak point or the like which allows a 1 portion of the container 16, 116 to be pulled or 2 twisted off. The sample could also be removed from the apparatus 10, 110 by providing on the second container 16, 116 a 'ring-pull' device or the like. 5 The sample could also be removed from the apparatus 7 10, 110 by providing on the second container 16, 116 a serrated cap portion or the like which is press-8 9 fitted onto the end portion 30, 130 of the container 10 The serrated cap portion is simply pulled 11 off when accessing the sample. The sample could also be removed from the apparatus 10, 110 by 12 13 providing on the second container 16, 116 a sliding 14 gate portion or the like which is simply slid into 15 an 'open' position when accessing the sample. 16 sample could also be removed from the apparatus 10, 110 by providing on the second container 16, 116 a 17 18 cap portion which may be pulled or slid into an 19 'open' position when accessing the sample. 20 sample could also be removed from the apparatus 10, 21 110 by providing on the second container 16, 116 a 22 rotating cap portion which allows one or more fluid 23 extraction points to be aligned with the second 24 chamber 18, 118 of the container 16, 116 when 25 accessing the sample.

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Although various methods have been described above for the removal of the sample from the apparatus 10, 11 for analysis, it should be appreciated that the sample may not necessarily need to be removed from the apparatus 10, 110 in order for the sample to be analysed. The apparatus 10, 110 may be constructed

of a material which is suitable for the sample to be

analysed whilst it is inside the second container

3 16, 116.

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5 Finally, although the first and second containers

6 12, 112 and 16, 116 have been described above as

7 being brought together by a machine which applies

8 the requisite force to end portions 28, 128 and 30,

9 130, it should be appreciated that the first and

second containers 12, 112 and 16, 116 could be

11 brought together by any other suitable method.